

1 I claim:

✓ 1. A method of treating a disease that results from a deficiency of a biological factor in a mammal wherein said method comprises administering Sertoli cells and a therapeutically effective amount of cells that produce said biological factor to a mammal in need of such treatment, wherein said Sertoli cells are administered in an amount effective to create an immunologically privileged site.

10 2. The method of Claim 1 wherein said mammal is a human.

3. The method of Claim 1 wherein said biological factor is a hormone.

15 4. The method of Claim 1 wherein said biological factor is insulin and said disease is diabetes mellitus.

5. The method of Claim 4 wherein said cells that produce said biological factor are pancreatic islet of Langerhans cells.

20 6. The method of Claim 1 wherein said cells that produce said biological factor are cells transformed by a nucleic acid encoding said biological factor.

25 7. The method of Claim 1 wherein said administering is by transplantation.

8. The method of Claim 1 wherein said Sertoli cells are administered in a dosage ranging from  $10^5$  to  $10^{10}$  cells.

30 9. The method of Claim 1 wherein said cells that produce said biological factor are administered in a dosage of from  $10^5$  to  $10^{10}$  cells.

35

08747122-110896

1           10. The method of Claim 7 wherein said  
transplantation is by xenograft.

          11. The method of Claim 7 wherein said  
transplantation is by allograft.

5           12. The method of Claim 1 which further  
comprises administering an immunosuppressive agent.

          13. The method of Claim 12 wherein said  
immunosuppressive agent is administered for a time  
sufficient to permit said transplanted cells to be  
10 functional.

          14. The method of Claim 12 wherein said  
immunosuppressive agent is cyclosporine.

          15. The method of Claim 14 wherein said  
cyclosporine is administered at a dosage of from 5 to 40  
15 mg/kg body wt.

          16. The method of Claim 1 which further  
comprises administering a therapeutically effective  
amount of exogenous biological factor following the  
transplantation of said cells that produce said  
20 biological factor.

          17. The method of Claim 1 wherein said cells  
that produce said biological factor are co-cultured with  
Sertoli cells in tissue culture.

          18. The method of Claim 17 wherein said cells  
that produce said biological factor are cryopreserved  
prior to co-culturing with Sertoli cells in tissue  
25 culture.

          19. A method of treating diabetes mellitus in  
a mammal wherein said method comprises administering to  
30 a diabetic mammal Sertoli cells in an amount effective  
to create an immunologically privileged site and a

1 therapeutically effective amount of pancreatic islet of Langerhans cells.

20. The method of Claim 19 wherein said diabetes mellitus is type I or type II.

5 21. The method of Claim 19 wherein said mammal is a human.

22. The method of Claim 19 wherein said Sertoli cells are human, bovine or porcine.

10 23. The method of Claim 19 wherein said pancreatic islet of Langerhans cells are human, bovine or porcine.

24. The method of Claim 19 wherein said administering is by transplantation.

15 25. The method of Claim 24 wherein said transplantation is by injection into the renal subcapsular space.

26. The method of Claim 24 wherein said transplantation is by injection into the subcutaneous facie.

20 27. The method of Claim 19 wherein said Sertoli cells are administered at a dosage ranging from  $10^5$  to  $10^{10}$  cells.

25 28. The method of Claim 19 wherein said islet of Langerhans cells are administered at a dosage ranging from 5-1000 islet cells/g body wt.

29. The method of Claim 19 which further comprises the administration of an immunosuppressive agent.

30 30. The method of Claim 29 wherein said immunosuppressive agent is administered for a time sufficient to permit the transplanted islets to be functional.

1           31. The method of Claim 29 wherein said  
immunosuppressive agent is cyclosporine.

5           32. The method of Claim 31 wherein said  
cyclosporine is administered at a dosage of 5 to 40  
mg/kg body wt.

10           33. The method of Claim 19 which further  
comprises administering a therapeutically effective  
amount of insulin following transplantation of said  
pancreatic islet of Langerhans cells.

10           ✓ 34. A method of creating an immunologically  
privileged site in a mammal wherein said method  
comprises transplanting isolated Sertoli cells into a  
mammal.

15           35. The method of Claim 34 wherein said mammal  
is a human.

36. The method of Claim 34 wherein said  
Sertoli cells are injected into the renal subcapsular  
space.

20           37. The method of Claim 34 wherein said  
Sertoli cells are injected into the subcutaneous facie.

38. The method of Claim 34 wherein said  
Sertoli cells are transplanted at a dosage ranging from  
 $10^5$  to  $10^{10}$  cells.

25           39. The method of Claim 34 wherein said  
Sertoli cells are human, bovine or porcine.

30           ✓ 40. A method of enhancing the recovery and  
proliferation of ex vivo cells comprising co-culturing  
said cells with Sertoli cells for a time and under  
conditions sufficient to achieve said enhanced recovery  
and proliferation.

101  
w/ 854

1        ✓ 41. A pharmaceutical composition comprising    161  
Sertoli cells and cells that produce a biological factor  
and a pharmaceutically acceptable carrier.

5        42. The composition of Claim 41 wherein said  
biological factor is a hormone.

43. The composition of Claim 41 wherein said  
cells that produce a biological factor are pancreatic  
islet of Langerhans cells.

10       44. The composition of Claim 41 wherein said  
cells that produce said biological factor are cells that  
are transformed by a nucleic acid encoding said  
biological factor.

15       ✓ 45. A pharmaceutical composition comprising  
Sertoli cells, pancreatic islet of Langerhans cells and  
a pharmaceutically acceptable carrier.

✓ 46. A pharmaceutical composition comprising  
Sertoli cells and a pharmaceutically acceptable carrier.

20       ✓ 47. A compartmentalized kit adapted to receive  
a first container adapted to contain Sertoli cells and a  
second container adapted to contain cells that produce a  
biological factor that is absent or defective in a  
disease.

25       ✓ 48. A compartmentalized kit adapted to receive  
a first container adapted to contain Sertoli cells and a  
second container adapted to contain pancreatic islet of  
Langerhans cells.

30       ✓ 49. An article of manufacture comprising a  
packaging material and Sertoli cells contained within  
said packaging material, wherein said Sertoli cells are  
effective for creating an immunologically privileged  
site in a mammal, and wherein said packaging material  
contains a label that indicates that said Sertoli cells

534  
NOV-D3V  
NOV-S17  
1534

1 can be used for creating an immunologically privileged  
site in a mammal.

/c

5

add  
B1

10

15

20

25

30

35

08747122-110896  
968077-2274780